

***IN THE UNITED STATES PATENT AND TRADEMARK OFFICE***

Applicant: Peng Cho TANG et al.  
Title: INDOLINONE DERIVATIVES AS PROTEIN  
KINASE/PHOSPHATASE INHIBITORS  
Prior Appl. No.: 09/871,700  
Prior Appl. Filing Date: 06/04/2001  
Examiner: Unassigned  
Art Unit: Unassigned

**INFORMATION DISCLOSURE STATEMENT**  
**UNDER 37 CFR §1.56**

Mail Stop PATENT APPLICATION  
Commissioner for Patents  
PO Box 1450  
Alexandria, Virginia 22313-1450

Sir:

Applicants submit herewith on Form PTO/SB/08 a listing of the documents cited by or submitted to the U.S. PTO in parent application Serial No. 09/871,700, filed 06/04/2001. As provided in 37 CFR §1.98(d), copies of the documents are not being provided since they were previously submitted to the United States Patent & Trademark Office in the above-identified parent application.

The submission of any document herewith, which is not a statutory bar, is not intended as an admission that such document constitutes prior art against the claims of the present application or that such document is considered material to patentability as defined in 37 CFR §1.56(b). Applicants do not waive any rights to take any action which would be appropriate to antedate or otherwise remove as a competent reference any document which is determined to be a *prima facie* art reference against the claims of the present application.

**TIMING OF THE DISCLOSURE**

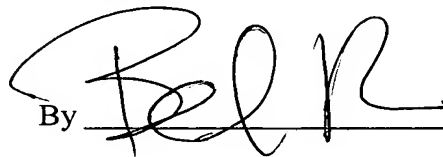
The listed documents are being submitted in compliance with 37 CFR §1.97(b), within three (3) months of the filing date of the application.

Applicants respectfully request that the listed document be considered by the Examiner and be made of record in the present application and that an initialed copy of Form PTO/SB/08 be returned in accordance with MPEP §609.

The Commissioner is hereby authorized to charge any additional fees which may be required regarding this application under 37 CFR §§ 1.16-1.17, or credit any overpayment, to Deposit Account No. 19-0741. Should no proper payment be enclosed herewith, as by a check being in the wrong amount, unsigned, post-dated, otherwise improper or informal or even entirely missing, the Commissioner is authorized to charge the unpaid amount to Deposit Account No. 19-0741.

Respectfully submitted,

Date 12/2/03

By 

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Customer Number: 22428

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Beth A. Burrous

Attorney for Applicant

Registration No. 35,087

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Substitute for form 1449B/PTO <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  <i>(use as many sheets as necessary)</i>				<b>Complete if Known</b>	
				<b>Prior Application Number</b>	09/871,700
				<b>Prior Appl. Filing Date</b>	06/04/2001
				<b>First Named Inventor</b>	Peng Cho TANG
				<b>Group Art Unit</b>	Unassigned
				<b>Examiner Name</b>	Unassigned
				<b>Attorney Docket Number</b>	034536-0907
Sheet	1	of	11		

U.S. PATENT DOCUMENTS						
Examiner Initials*	Cite No. <sup>1</sup>	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number	Kind Code <sup>2</sup> (if known)			
	A1	2,968,557		Burgandt et al.	01-17-1961	
	A2	4,002,749		Rovnyak	01-11-1977	
	A3	4,053,613		Rovnyak et al.	10-11-1977	
	A4	4,642,309		Michel et al.	02-10-1987	
	A5	4,826,847		Michel et al.	05-02-1989	
	A6	5,051,417		Nadler et al.	09-24-1991	
	A7	5,124,347	A	Connor et al.	06-23-1992	
	A8	5,196,446	A	Levitzi et al.	03-23-1993	
	A9	5,302,606	A	Spada et al.	04-12-1994	
	A10	5,322,950	A	Sircar et al.	06-21-1994	
	A11	5,374,652	A	Buzzetti et al.	12-20-1994	
	A12	5,382,593	A	Le Baut et al.	01-17-1995	
	A13	5,389,661	A	Sircar et al.	02-14-1995	
	A14	5,397,787	A	Buzzetti et al.	03-14-1995	
	A15	5,409,949	A	Buzzetti et al.	04-25-1995	
	A16	5,792,783	A	Tang et al.	08-11-1998	
	A17	5,834,504	A	Tang et al.	11-10-1998	
	A18	5,849,710	A	Battistini et al.	12-15-1998	
	A19	5,880,141	A	Tang et al.	03-09-1999	

Examiner Signature	Date Considered
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<sup>5</sup> Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup> Applicant is to place a check mark here if English language Translation is attached.

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	A20	5,883,113	A	Tang et al.	03-16-1999	
	A21	5,883,116	A	Tang et al.	03-16-1999	
	A22	5,886,020	A	Tang et al.	03-23-1999	
	A23	6,133,305	A	Tang et al.	10-17-2000	

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	A24	WO	91/13055	A2	FARMITALIA CARLO ERBA SRL	09-05-1991		
	A25	WO	92/07830	A2	PFIZER INC.	05-14-1992		
	A26	WO	92/20642	A1	RHONEPOULENC RORER INTERNATIONAL	11-26-1992		
	A27	WO	93/01182	A1	FARMITALIA CARLO ERA SRL	01-21-1993		
	A28	WO	94/14808	A1	FARMITALIA CARLO ERBA SRL	07-07-1994		
	A29	WO	95/01349	A1	FARMITALIA CARLO ERBA SRL	01-12-1995		
	A30	WO	95/17181	A1	PHARMACIA S.P.A.	06-29-1995		
	A31	WO	96/00226	A1	PHARMACIA S.P.A.	01-04-1996		
	A32	WO	96/16964	A1	PHARMACIA S.P.A.	06-06-1996		
	A33	WO	96/22976	A1	PHARMACIA S.P.A.	08-01-1996		

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	A34	WO	96/32380	A1	PHARMACIA S.P.A.	10-17-1996		
	A35	WO	96/40116	A1	SUGEN, INC.	12-19-1996		
	A36	WO	97/25986	A1	TAIHO PHARMACEUTICAL CO., LTD.	07-24-1997		
	A37	WO	98/07695	A1	SUGEN, INC.	02-26-1998		
	A38	WO	98/24432	A2	SUGEN, INC.	06-11-1998		
	A39	WO	98/38984	A2	SUGEN, INC.	09-11-1998		
	A40	WO	98/50356	A1	SUGEN, INC.	11-12-1998		
	A41	WO	98/50356	A	SUGEN, INC.	11-12-1998		
	A42	WO	99/10325	A1	GLAXO GROUP LIMITED	03-04-1999		
	A43	WO	99/52869	A1	BOEHRINGER INGELHEIM PHARMA KG	10-21-1999		
	A44	WO	99/61422	A1	SUGEN, INC.	12-02-1999		
	A45	WO	99/65869	A1	BAYER AKTIENGESELLSCHAFT	12-23-1999		
	A46	WO	00/08202	A2	SUGEN, INC.	02-17-2000		
	A47	WO	00/35906	A2	F. HOFFMAN – LA ROCHE AG	06-22-2000		
	A48	WO	00/35908	A1	F. HOFFMAN – LA ROCHE AG	06-22-2000		
	A49	WO	00/35909	A1	F. HOFFMAN – LA ROCHE AG	06-22-2000		
	A50	WO	00/56709	A1	SUGEN, INC.	09-28-2000		

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	A51	WO	01/60814	A2	SUGEN, INC.	08-23-2001		
	A52	WO	02/02551	A	SUGEN, INC.	01-10-2002		
	A53	DE	2,159,360	A	BAYER AG	06-14-1973		X
	A54	DE	2,159,361	A	BAYER AG	06-14-1973		X
	A55	DE	2,159,362		BAYER AG	06-14-1973		
	A56	DE	2,159,363	A	BAYER AG	06-14-1973		X
	A57	DE	2,321,656	A	COLGATE-PALMOLIVE CO.	11-15-1973		X
	A58	DE	3,426,419	A	BOEHRINGER MANNHEIM GMBH	01-23-1986		X
	A59	EP	0 252 713	B1	PFIZER INC.	01-13-1988		
	A60	EP	0 351 213	A2	LES LABORATOIRES BEECHAM S.A.	01-17-1990		
	A61	EP	0 525 472	A2	FARMITALIA CARLO ERBA SRL	02-03-1993		
	A62	EP	0 632 102	A1	BAYER AG	01-04-1995		X
	A63	EP	0 662 473	A1	PHARMACIA S.P.A.	07-12-1995		
	A64	EP	0 769 947	B1	TANG, Peng Cho et al.	05-02-1997		
	A65	EP	0 788 890	A1	AGFA-GEVAERT	08-13-1997		
	A66	EP	0 934 931	A2	SUGEN, INC.	08-11-1999		
	A67	EP	1 082 305	A1	SUGEN, INC.	03-14-2001		
	A68	FR	1.398.224		IMPERIAL CHEMICAL INDUSTRIES LIMITED	05-07-1965		X

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		Office <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	A69	FR	1.599.772		INSTITUT PASTEUR	08-28-1970		X
	A70	FR	2.689.397	A1	ADIR ET COMPAGNIE	10-08-1993		X
	A71	GB	809,691		Roy HULL	03-04-1959		
	A72	GB	835,473		Norman SENIOR	05-18-1960		
	A73	JP	62-29570	A	KANEGAFUCHI CHEM KK	02-07-1987		X
	A74	JP	62-39564	A	KANEGAFUCHI CHEM KK	02-20-1987		X
	A75	JP	63-141955	A	KANEGAFUCHI CHEM KK	06-14-1988		X
	A76	JP	5-58894	A	KANEKA CORP	03-09-1993		X
	A77	CA	2,012,634	A1	UNIVERSITY OF BRITISH COLUMBIA	09-20-1991		
	A78	AU	286870		IMPERIAL CHEMICAL INDUSTRIES OF AUSTRALIA AND NEW ZEALAND LIMITED	05-11-1967		

OTHER PRIOR ART – NON PATENT LITERATURE DOCUMENTS				
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>6</sup>	
	A79	ANDREANI et al., "Potential Antitumor Agents. 25[1]. Synthesis and Cytotoxic Activity of 3-(2-Chloro-3-Indolymethylene)1,3-Dihydroindol-2-Ones," <u>Anticancer Research</u> 16:3585-3588 (1996) © Elsevier, Paris		
	A80	ANDREANI et al., "Synthesis and cardiotoxic activity of 2-indolinones," <u>Eur. J. Med. Chem.</u> 25:187-190 (1990)		
	A81	ANDREANI et al., "Synthesis and cardiotoxic activity of 2-indolinones bearing pyridyl groups," <u>Eur. J. Med. Chem.</u> 28:653-657 (1993) © Elsevier, Paris		

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	A82	ANDREANI et al., "Synthesis and cardiotonic activity of pyridylmethylene-2-indolinones," <u>Eur. J. Med. Chem.</u> 27:167-170 (1992) © Elsevier, Paris		
	A83	ANDREANI et al., "Synthesis and potential coanthracyclic activity of substituted 3-(5-imidazo[2,1-b]thiazolylmethylene)-2-indolinones," <u>Eur. J. Med. Chem.</u> 32:919-924 (1997) © Elsevier, Paris		
	A84	ANDREANI et al., "Synthesis of lactams with potential cardiotonic activity," <u>Eur. J. Med. Chem.</u> 28:825-829 (1993)		
	A85	ANDREANI et al., "In Vivo Cardiotonic Activity of Pyridylmethylene-2-indolinones," <u>Arzneimittel-Forschung Drug Research</u> 48:727-729 (1998) ©		
	A86	BAHNER and BROTHERTON, "6-Dimethylaminochrysene and Other Analogs of 4-(4-Dimethylamino)stilbene," <u>J. Med. Chem.</u> 12:722-723 (1969)		
	A87	BAHNER et al., "Benzylideneindenes with Oxygen Attached to the Indene Ring," <u>J. Med. Chem.</u> 12:721-722 (1969)		
	A88	BAMFIELD et al., "Diels-Alder Reactions of Oxindolylideneacetone," <u>J. Chem. Soc. (C)</u> 1028-1030 (1966) ©		
	A89	BORSCHKE et al., "Über vielkernige kondensierte Systeme mit heterocyclischen Ringen. XIII.," <u>Liebigs Ann. Chem.</u> 550:160-174 (1941)		
	A90	BUZZETTI et al., "Cinnamamide Analogs as Inhibitors of Protein Tyrosine Kinases," <u>Il Farmaco</u> 48:615-636 (1993)		

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	A91	CHATTEN et al., "Substituted Oxindoles. Part VI. Polarographic Reduction of Substituted <i>trans</i> -3-Benzylideneindol-2(3 <i>H</i> )-ones," <u>J. Chem. Soc. Perkin II</u> : 469-473 (1973)	
	A92	CODA et al., "(Z)- and (E)-Arylidene-1,3-dihydroindol-2-ones: Configuration, Conformation and Infrared Carbonyl Stretching Frequencies," <u>J. Chem. Soc. Perkin Trans. II</u> : 615-619 (1984)	
	A93	CODA et al., "3-(4-methylbenzylidene)-1,3-dihydroindol-2-one," <u>Journal of the Chemical Society, Perkin Transactions 2</u> 4:615-620 (1984) DATABASE CROSSFIRE, Beilstein Reference No. 6-21	
	A94	DECODTS et al., "Suicide inhibitors of proteases. Lack of activity of halomethyl derivatives of some aromatic lactams," <u>Eur. J. Med. Chem</u> 18: 107-111 (1983)	
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	A98	GAZIT et al., "Tyrphostins. 2. Heterocyclic and $\alpha$ -Substituted Benzylidenemalononitrile Tyrphostins as Potent Inhibitors of EGF Receptor and ErbB2/neu Tyrosine Kinases," <u>J. Med. Chem.</u> 34:1896-1907 (1991) copyright Am. Chem. Soc.	
	A99	HIRAO et al., "Rhodium-Catalyzed Carbonylation of 2-Alkynylaniline: Syntheses of 1,3-Dihydroindol-2-ones," <u>Tetrahedron Letters</u> 36(35) 1995 ©Pergamon	

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				Prior Appl. Filing Date	06/04/2001
				First Named Inventor	Peng Cho TANG
				Group Art Unit	Unassigned
				Examiner Name	Unassigned
Sheet	8	of	11	Attorney Docket Number	034536-0907

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	A100	HODGES et al., "Chemical and biological properties of some oxindolidyl-3-methines," <u>Canadian J. Chemistry</u> 46:2189-2194 (1968)		
	A101	HOWARD, Harry R., "Lactam Derivatives," U.S. Provisional Patent Application Number 60/015134		
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	A103	KATRITZKY et al., "Color and Constitution. Part 8[1]. Some Novel Dyestuffs Containing Indoxyl Residues," <u>J. Heterocyclic Chem.</u> 25:1287-1292 (1988)		
	A104	KOBAYASHI et al., "Anti-tumor Activity of Indole Derivatives," <u>Yakugaku Zasshi</u> 97:1033-1039 (1977)		
	A105	KOVAC and STETINOVA, "Furan derivatives. LXXX. Synthesis and properties of substituted furfurylidenoxindoles," <u>Chem. rvesu</u> 30:484-492 (1976)		
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	A107	MARIANI et al., "Inhibition of angiogenesis by FCE 26806, a potent tyrosine kinase inhibitor," <u>Experimental Therapeutics - Proceedings of the American Association for Cancer Research</u> 35:381 at abstract no. 2268 (March 1994)		
	A108	MOHAMMADI et al., "Structures of the Tyrosine Kinase Domain of Fibroblast Growth Factor Receptor in Complex with Inhibitors," <u>Science</u> 276:955-960 (1997) © American Association for the Advancement of Science		

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	A109	NEBER and RÖCKER, "On the action of benzaldehydes on the free o-aminophenylacetic acid (II)," <u>Chem. Ber.</u> 56:1710-1716 (1923) (GERMAN AND ENGLISH TRANSLATION)		
	A110	NODIFF et al., "Antimalarial Phenanthrene Amino Alcohols. 3. Halogen-containing 9-phenanthrenemethanols," <u>Chemical Abstracts</u> , Vol. 83, abstract no. 188214 (1975)		
	A111	O'SULLIVAN and ROTHERY, "The Preparation and Possible Clinical Significance of 4'-Dialkylaminoisoindogenides," <u>Clinica Chimica Acta</u> 62:181-182 (1975) ©Elsevier Scientific Publishing Company		
	A112	PAVLENKO et al., "Introduction of aminomethyl groups into heterocyclic CH-acid molecules," <u>Dopov. Akad. Nauk Ukr Rsr.</u> , Ser. B: Geol., Khim. Biol. Nauki 7:64-66 (1980) We should add that we are Sub. Abstract		
	A113	PLOWMAN et al., "Receptor Tyrosine Kinases as Targets for Drug Intervention," <u>DN&amp;P</u> 7:334-339 (1994)		
	A114	QUALLICH et al., "A General Oxindole Synthesis," <u>J. Synthetic Organic Chemistry</u> : 51-51 (1993)		
	A115	SCHUCHTER et al., "Successful Treatment of Murine Melanoma with Bryostatin 1," <u>Cancer Research</u> 51:682-687 (1991)		
	A116	SHIRAISHI et al., "Specific inhibitors of Tyrosine-Specific Protein Kinase, Synthetic 4-Hydroxycinnamamide Derivatives," <u>Biochemical and Biophysical Research Communications</u> 147:322-328 (1987)© Academic Press		
	A117	SHIRAISHI et al., "Specific Inhibitors of Tyrosine-specific Protein Kinases: Properties of 4-Hydroxycinnamamide Derivatives <u>in Vitro</u> ," <u>Cancer Research</u> 49:2374-2378 (1989)		

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	A118	SINGH et al., "Indolinone Derivatives as Potential Antimicrobial Agents," <u>Zentralbl. Mikrobiol.</u> 144:105-109 (1989) copyright VEB Gustav Fischer Verlag Jena		
	A119	SPADA, et al., "Small molecule inhibitors of tyrosine kinase activity," <u>Expert Opinion on Therapeutic Patents</u> 5:805-817 (1995) ©Ashley Publications		
	A120	SUN et al., "Design, Synthesis, and Evaluations of Substituted 3-[(3- or 4-Carboxyethylpyrrol-2-yl) methylidene]indolin-2-ones as Inhibitors of VEGF, FGF, and PDGF Receptor Tyrosine Kinases," <u>Journal of Medicinal Chemistry</u> 42: 5120-5130 (1999) ©American Chemical Society		
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	A122	TACCONI and MARINONE, "Preparazione e caratteristiche di alcuni 3-ossindolidenderivati," <u>Ricerca Scientifica</u> 38:1239-1244 (1968)		
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	A125	TRAXLER, "Protein tyrosine kinase inhibitors in cancer treatment," <u>Expert Opinion on Therapeutic Patents</u> 7(6):571-588 (1997) © Ashley Publications Ltd.		
	A126	von DOBENECK, H. et al., "α-β'-Diindolylmethane und –methene. Der Urorosein-Chromophor," <u>Chemische Berichte</u> , 102(4):1347-1356 (1969)		

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	A133	ZHANG et al., "Microtubule Effects of Welwistatin, a Cyanobacterial Indolinone that Circumvents Multiple Drug Resistance," <u>Molecular Pharmacology</u> 49:228-234 (1996) ©The American Society for Pharmacology and Experimental Pharmaceutics		
	A134	ZHUNGIETU et al., "Reaction of Indoles and 2-Ketoindolines With Some Aldehydes," <u>Chemical Abstracts</u> , Vol. 78, abstract no. 111201 (1990)		

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